MCP-IRA

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GenCore version 5.1.4_p5_4578
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OM protein - protein search, using sw model

Run on: May 19, 2003, 16:35:42; Search time 55.5395 Seconds
(without alignments)
897.302 Million cell updates/sec

Title: US-09-625-573-2
Perfect score: 1970
Sequence: 1 MLSTSRSFFIRNTNESGEEV......GKGKSIGRAPEASLQDKEGA 374

Scoring table: BLOSUM62 Gapop 10.0 , Gapext 0.5

Searched: 908470 seqs, 133250620 residues total number of hits satisfying chosen parameters: 9

Minimum DB seq length: 0
Maximum DB seq length: 200000000

Post-processing: Minimum Match 0% Maximum Match 100% Listing first 45 summaries

Database : A

A_Geneseq_101002:*

| SIDS2/gcgdata/geneseq/genesegp-embl/AA1980.DAT:*
| SIDS2/gcgdata/geneseq/genesegp-embl/AA1981.DAT:*
| SIDS2/gcgdata/geneseq/genesegp-embl/AA1982.DAT:*
| SIDS2/gcgdata/geneseq/genesegp-embl/AA1983.DAT:*
| SIDS2/gcgdata/geneseq/genesegp-embl/AA1983.DAT:*
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| SIDS2/gcgdata/geneseq/genesegp-embl/AA1999.DAT:*
| SIDS2/gcgdata/geneseq-embl/AA1999.DAT:*
| SIDS2/gcgdata/gen

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

Description	Human monocyte che	Human CCR2a protei	Human MCP-1 recept	Human monocyte che	Human monocyte che	Human CCR2b protei	Human wild-type CC	Human CCR2-64I pol	Non-endogenous hum	Amino acid sequenc
ID	AAR79165	AAG80107	AAB46859	AAR79166	AAW35833	AAG80108	AAU07614	AAU07613	ABB56340	. AAG79089
DB	16	22	22	16	18	22	22	22	22	22
% Query Match Length DB	. 374	374	329	360	360	360	360	360	360	352
% Query Match	100.0	100.0	87.7	83.8	83.8	83.8	83.8	83.8	83.5	62.7
Score	1970	1970	1727.5	1651.5	1651.5	1651.5	1651.5	1650.5	1645.5	1236
Result No.	Т	7	e	4	5	9	7	80	6	10

Mouse CC-CKR5 prot Human CCR5. Homo Human chemokine re Macaque chemokine Human CCR5 protein Human CCR5 protein	rotei rotei pro nokin NR10 rotei nokin	Human CC chemokine Non endogenous hum Human C-protein ch Human G-protein ch	C-C chemokine rece Human MIP-1 alpha/R Human MIP-1alpha/R Human CC-chemokine Rat CC chemokine r Human C-C chemokin CC-chemokine recep Human C-C chemokin Human C-C chemokin
AAW54037 AAW27123 AAW27125 AAW27125 AAW88232 AAW88232	AAB82948 AAE07039 AAE07039 AAB93354 AAB04321 AAB46858 AAB97152 ABB08343 AAB52828	0 M 4 M O M M 4 M O .	AAK25749 AAW2578 AAW2571 AAW29179 AAW3174 AAW3174 AAW10100
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113 114 115 117	20000000000000000000000000000000000000	332 332 343 36 36 37 37	78883338 0044444 0178445
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ALIGNMENTS

Monocyte chemoattractant protein-1 receptor; MCR-1R; chemokine. Human monocyte chemoattractant protein-1 receptor MCP-1RA. 49..70 /label= transmembrane 80..700 204..231 /label= transmembrane 1..48 /label= extracellular /label= transmembrane transmembrane 'label= transmembrane 'label transmembrane /label= transmembrane /label= carboxyl tail Location/Qualifiers AAR79165 standard; Protein; 374 AA. 29-DEC-1995 (first entry) ..313 .178 244..268 115..136 314.375 label= Homo sapiens AAR79165; Key Domain Domain Domain Domain Domain Domain Domain Domain Region AAR79165

95WO-US00476 94US-0182962.

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family, degenerate oligo primers were designed corresp. to the conserved sequences R79167 in the second and R79168 in the third cransmembrane domains of the MIP-lalpha/RANTES receptor, the IL-8 receptors and the HUMSTRS orphan receptor (GenBank Accession #M9923. The degenerate oligo incorporating EcoRI and XhoI sites at their 5. Cends are 096299 and 096300. Amplification of cDNA derived from MM6 cells with the primers yieled a number of cDNA derived from MM6 cells with the primers yieled a number of ECORI and Ill-length version of this clone, a MM6 cDNA library was constructed in pFRGO and probed with the PCR product. A 2.1 kb cDNA clone was obtd. Analysis of additional clones in the MM6 cDNA library revealed a second sequence that was identical to the 2.1 kb cDNA sequence first obtd. From the 5' UTR through the puttive seventh transmembrane domain but contained a different cytoplasmic tail. The second sequence appears to represent alternative splicing of the carboxyl-terminal conditions the MCP-IR protein. The two sequences are denoted MCP-IRA and MCP-IRA as and MCP-IRB (see 096297/R79165 & 096298/R79166). Active mature MAS a moll wt of about 42,000 daltons. MCP-IRB
                                                                                                                                                                                                                                                                                                                                                  DNA encoding monocyte chemo-attractant protein-1 receptor - used pa
for identifying antagonists and for treating diseases characterised
monocytic infiltrates
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         To identify and clone new members of the chemokine receptor gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   MCP-IRA has a predicted mol. wt. of abc
has a mol. wt. of about 41,000 daltons.
                                                                                                                                                                                                                                                                                                                                                                                                                                     Claim 2; Fig 1; 84pp; English.
                                                                                                                                                                                             (REGC ) UNIV CALIFORNIAL.
                                                                                                                                                                                                                                         Charo I, Coughlin S;
                                                                                                                                                                                                                                                                                  WPI; 1995-263866/34.
                                                                                                                                                                                                                                                                                                         N-PSDB; AAQ96297
                                                                                                         11-JAN-1995;
                                                                                                                                                  13-JAN-1994;
                  W09519436-A.
                                                             20-JUL-1995.
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374 AA; Sequence

Gaps 0; 100.0%; Score 1970; DB 16; Length 374; 100.0%; Pred. No. 2.1e-216; Live 0; Mismatches 0; Indels 0; 100.0%; L.v. 0, Conservative Best Local Similarity Matches 374; Query Match

; 0

61 MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120 1 MLSTSRSRFIRNTNESGEEVTTFFDYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGN 60 g δ g

121 HIGYFGGIFFIILLTIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTK 180 CQKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR 240 181 δ g ò

241 AVRVIFTIMIVYFLEWTPYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCI 300 δ a

181

QQ

NPIIYAFVGEKFRSLFHIALGCRIAPLQKPVCGGPGVRPGKNVKVTTQGLLDGRGKGKSI 360 301 qq ò

Diagnostic agent containing two or more receptor-specific ligands, useful for detecting tumors, inflammation etc., also therapeutic use of ligand inhibitors Chemokine; tumour diagnosis; colorectal; prostatic; organ rejection; inflammation; autoimmune disease; metastasis; bronchial asthma; lupus; chronic bowel inflammation; rheumatoid arthritis; cytostatic; antiinflammatory; antiasthmatic; immunosuppressive; dermatological; antirheumatic; antiarthritic. Heitland A, Spodsberg N; AAG80107 standard; Protein; 374 AA. Disclosure; Page 9; 26pp; German. 02-APR-2001; 2001WO-EP03708. 31-MAR-2000; 2000DE-1016013. (first entry) Forssmann W, Adermann K, 361 GRAPEASLQDKEGA 374 361 GRAPEASLQDKEGA 374 (IPFP-) IPF PHARM GMBH. Human CCR2a protein. WPI; 2001-626256/72. (FORS/) FORSSMANN U. WO200172830-A2. Homo sapiens, 17-JAN-2002 04-OCT-2001. AAG80107; RESULT 2 AAG80107 δλ Db

partic.

This invention describes a novel diagnostic agent (A) comprising at least two different ligands (I) for receptors (II) that are implicated in disease. (A) are used for the diagnosis of tumors (especially colorectal or prostatic), organ rejection, inflammation and autoimmune diseases. Also inhibitors of (I) are used therapeutically against tumors (and their metastases), inflammation, particularly bronchial asthma or chronic bowel inflammation), or autoimmune diseases (rheumatodia arthritis or lupus), where the (cardio)vascular, lymphatic, respiratory, nervous, digestive, andocrine, motor or urogenital systems or skin are affected, and bone marrow diseases. The products of the invention are chemokine derivatives which have cytostatic, antiinflammatory, antiasthmatic, antiinflammatory, antiasthmatic, chemokines act on specific tumor and inflammatory calls through a constellation of chemokine receptors (CR), which constellation of these cells. AAG80128 represent human chemokine fragments used to illustrate the method of the invention.

374 AA; Seguence

ö Gaps 1 MLSTSRSRFIRNTNESGEEVTTFFDYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGN 60 ; 100.0%; Score 1970; DB 22; Length 374; 100.0%; Pred. No. 2.1e-216; Indels Mismatches 0; Conservative Query Match Best Local Similarity Matches 374; δλ

qq

61 MLVVLILINCKKLKCLTDIYLENLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120 δλ

Qλ q δ

Óγ g δ

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Domain
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               HDGNR10; human; G-protein chemokine receptor; antiinflammatory; immunomodulatory; anticoagulant; antiallergic; immunosuppressive; cytostatic; antiparastic; antiparastic; antiparastic; antiparastic; antiparastic; anticonatic; and the subject of the second of the second
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     This invention describes a novel receptor polypeptide (I) selected from (i) a fully defined 329 amino acid sequence (II) fully disclosed in the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      New human G-protein chemokine receptor polypeptides and polynucleotides, useful for identifying (ant)agonists to the G-protein chemokine receptor -
                                                                                                                                                                                                                                                                            61 MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLFLWAHSAANEWVFGNAMCKLFTGLY 120
                                     HIGYFGGIFFIILLTIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTK 180
                                                                                                           CQKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR 240
                                                                                                                                                                                                                                                           NPIIYAFVGEKFRSLFHIALGCRIAPLQKPVCGGPGVRPGKNVKVTTQGLLDGRGKGKSI 360
                                                      AVRVIFTIMIVYFLFWTPYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Disclosure; Page 16-17; 22pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Human MCP-1 receptor protein fragment.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AAB46859 standard; Protein; 329 AA.
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98US-0195662.
99US-0339912.
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                                                                                                                                                                                                                                                                                                                                                            361 GRAPEASLODKEGA 374
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(RUBE/) RUBEN S M.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Ruben SM;
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18-NOV-1998;
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plasmid, and fragments, analogs and derivatives of the polypeptide. The plasmid, and fragments, analogs and derivatives of the polypeptide. The products of the invention have antiliflammatory, immunomodulatory, anticoagulant, antiallergic, immunosuppressive, vulnerary, cytostatic, antiparasitic, antipsoriatic, antirheumatic, antiarthritic and vasotropic activity and can be used for gene therapy. The G-protein chemokine receptors, HDGNRIO, (I) are useful for screening for compounds which activate or inhibit activation of (I). The products of the invention can also be used for stimulating heematopolesis, wound healing, coagulation, angiogenesis, treating solid tumours, chronic infections, leukemia, carimulating growth factor activity. HDGNRIO is useful for treating callergy, atherogenesis, anaphylaxis, malignancy, chronic and acute inflammation, histanhae and immunoglobulin E (IgE)-mediated allergic reactions, prostaglandin independent fever, bone marrow failure, sallicosis, sarcoidosis, rheumatoid arthritis, shock and hyper-ecsimophilic syndrome.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLYHIGYFGGIFFIILLTID 137
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human monocyte chemoattractant protein-1 receptor MCP-1RB.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               87.7%; Score 1727.5; DB 2:
95.6%; Pred. No. 9.7e-189;
Live 0; Mismatches 0;
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80..700
/label= transmembrane
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Matches 329; Conserv
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ij

244..268 /label= transmembrane label= transmembrane

/label= transmembrane

..231

Domain Domain Domain Region Domain

295..313 /label= transmembrane

1..48 /label= extracellular 314..360 /label= carboxyl

> WO9519436-A. 20-JUL-1995

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95WO-US00476
                                                                                               94US-0182962
                                                                                                         (REGC ) UNIV CALIFORNIA.
                                                                                                                                                          monocytic infiltrates
                                                                                                                    Charo I, Coughlin S;
                                                                                                                              WPI; 1995-263866/34.
                                                                                                                                    N-PSDB; AAQ96298
                                                                                    11-JAN-1995;
                                                                                               13-JAN-1994;
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- used partic. DNA encoding monocyte chemo-attractant protein-1 receptor - used part: for identifying antagonists and for treating diseases characterised by

Claim 2; Fig 2; 84pp; English.

conserved sequences R79167 in the second and R79168 in the third transmembrane domains of the MIP-lalpha/RANTES receptor, the LI-8 transmembrane domains of the MIP-lalpha/RANTES receptor, the LI-8 receptors and the HUMSTRS orphan receptor (GenBank Accession #M99293.

The degenerate oligo incorporating EcoRI and XhoI sites at their 5 cends are 0965299 and 096300. Amplification of cDNA derived from MM6 and are 0965299 and 096300. Amplification of cDNA derived from MM6 cells with the primers yieled a number of PCR products. One cDNA appeared to encode a novel protein. To obtain a full-length version of this clone, a MM6 cDNA library was constructed in PFRGG and probed with the PCR product. A 2.1 kb cDNA clone was obtd. Analysis of additional clones in the MM6 cDNA library revealed a second sequence that was identical to the 2.1 kb cDNA sequence first obtd. Crome the 5' UTR through the putative seventh transmembrane domain but contained a different cytoplasmic tail. The second sequence appears to represent alternative splicing of the carboxyl-terminal tail of the MCP-IRA product and MCP-IRA has a predicted mol. Wt. of about 42,000 daltons. MCP-IRB conditions. MCP-IRB conditions. identify and clone new members of the chemokine receptor gene

360 AA; Seguence

SS X 8

Query Match

MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120 Gaps 1 MLSTSRSFIRNTNESGEEVTTFFDYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGN 60 83.8%; Score 1651.5; DB 16; Length 360; 95.5%; Pred. No. 5.4e-180; ive 3; Mismatches 5; Indels 7; Best Local Similarity 95.5 Matches 319; Conservative 61 g òλ ò

CQKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR 240 181

61

q ò ОР 181 CQKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR 240

ΩD ò

The present sequence represents human monocyte chemoattractant protein 1 (MCP-1) receptor protein. The MCP-1 receptor protein and encoding DNA are used for the prevention and treatment of tumours and inflammatory, viral, infectious, allergic, diabetic and central nervous system DNA encoding human monocyte chemoattractant protein 1 receptor - used to treat tumours and inflammatory, viral, infectious, allergic, diabetic and central nervous system diseases monocyte chemoattractant protein; receptor; tumour; į Human monocyte chemoattractant protein 1 receptor. inflammatory disease; viral; allergy; diabetes 301 NPIIYAFVGEKFR---SLF---HIALG-CRIAPL 327 301 NPIIYAFVGEKFRRYLSVFFRKHITKRFCKQCPV 334 Disclosure; Page 12-14; 15pp; Japanese. AA. AAW35833 standard; Protein; 360 (TAKE) TAKEDA CHEM IND LTD. 96JP-0053574 96JP-0053574 27-FEB-1998 (first entry) WPI; 1997-506557/47 360 AA; N-PSDB; AAT96976. Human; MCP-1; JP09238688-A. 11-MAR-1996; Homo sapiens 11-MAR-1996; 16-SEP-1997. AAW35833; Sequence diseases Query Match RESULT 5 AAW35833 ŏ qq δy q

61 MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120 HIGYFGGIFFIILLTIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTK 180 COKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR 240 Gaps 1 MLSTSRSRFIRNTNESGEEVTTFFDYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGN 83.8%; Score 1651.5; DB 18; Length 360; 95.5%; Pred. No. 5.4e-180; 7; Indels 3; Mismatches Conservative Best Local Similarity Matches 319; Conserv 121 181

g δλ QQ δy qq

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a δλ Д

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This invention describes a novel diagnostic agent (A) comprising at least two different ligands (I) for receptors (II) that are implicated in disease. (A) are used for the diagnosis of tumors (especially colorectal or prostatic), organ rejection, inflammation and autoimmune diseases. Also inhibitors of (I) are used therapeutically against tumors (and their metastases), inflammation (particularly bronchial asthma or chronic bowel inflammation), where the (cardio)vascular, Impunbatic, respiratory, nervous, digestive, endocrine, motor or urogenital systems or skin are affected, and bone marrow diseases. The products of the invention are chemokine derivatives which have contracted to the invention are chemokine derivatives
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      which have cytostatic, antiinflammatory, antiasthmatic, immunosuppressive, dermatological, antirheumatic, antiarthritic. Chemokines act on specific tumor and inflammatory cells through a constellation of chemokine receptors (CR), which control migration and proliferation of these cells. AAG80045-AAG80128 represent human chemokine fragments used to illustrate the method of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            useful for detecting tumors, inflammation etc., also therapeutic use of ligand inhibitors
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    inflammation; autoimmune disease; metastasis; bronchial astimu; lupus; chronic bowel inflammation; rheumatoid arthritis; cytostatic; antiinflammatory; antiasthmatic; immunosuppressive; dermatological; antirheumatic; antiarthritic.
                                  241 AVRVIFTIMIVYFLEWTPXNIVILLINTEQEFFGLSNCESTSQLDQATQVTETLGMTHCCI 300
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Chemokine; tumour diagnosis; colorectal; prostatic; organ rejection;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     or more receptor-specific ligands
241 AVRVIETIMIVYFLFWTPYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMIHCCI
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                                                                                                                                                                                                 301 NPIIYAFVGEKFR---SLF---HIALG-CRIAPL 327
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                                                                                                                                                                                                                                                                                                                                                                                                              AAG80108 standard; Protein; 360 AA
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             17-JAN-2002 (first entry)
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             (IPFP-) IPF PHARM GMBH. (FORS/) FORSSMANN U.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human CCR2b protein.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 2001-626256/72.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        04-OCT-2001.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AAG80108;
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                                                                                                                                                                                                                                                                                                                                                                               AAG80108
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The invention relates to diagnosing atherosclerosis (or susceptibility to) in a subject by determining expression or activity of the human CCR2-64I polymeptide (a polymorphic variant form of the human CCR2 receptor) or the CCR2-64V polymeptide (human CCR2 receptor), by screening for a single nucleotide polymorphism in codon 64 of the polynucleotide encoding the CCR2 receptor. This results in production of CCR2-64I, whereby polymorphic variants are associated with a lower incidence of atherosclerosis. The presence or amount of CCR2-64IV in a sample can also be analysed. The sequences of the invention can be used for predicting the response of a patient to drug treatment, for predicting the disease outcome in a patient and also for the production of a treatment for hypercholesterolaemia. The sequence represents the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Human; CCR2 receptor; CCR2-641; CCR2-64V; gene therapy; atherosclerosis; single nucleotide polymorphism; hypercholesterolaemia.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          subject, by determining a single nucleotide polymorphism in specific codon of a polynucleotide encoding human CCR2 receptor in genome of the
                                                                                                                                                                                                                                                                                    241 AVRVIETIMIVYFLEWTPYNIVILLINTEQEFFGLSNCESTSQLDQATQVTETLGMTHCCI 300
                                                                                                                                                                                                                                                                  241 AVRVIFTIMIVYFLEWTPYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCI 300
                                                                                                                         HIGYFGGIFFIILLTIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTK 180
                                                     MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120
                                                                                      MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Diagnosing atherosclerosis or susceptibility to atherosclerosis in a
COKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR
                                                                                                                                                                                                                                                                                                                                        NPIIYAFVGEKFR---SLF---HIALG-CRIAPL 327
                                                                                                                                                                                                                                                                                                                                                              Human wild-type CCR2-64V polypeptide.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AAU07614 standard; Protein; 360 AA.
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AAU07614
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5; Indels

DB 22; Length 360;

83.8%; Score 1651.5; DB 2 95.5%; Pred. No. 5.4e-180;

3; Mismatches

Matches 319; Conservative

Local Similarity

Query Match

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Human; CCR2 receptor; CCR2-641; CCR2-64V; gene therapy; atherosclerosis; single nucleotide polymorphism; hypercholesterolaemia.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Diagnosing atherosclerosis or susceptibility to atherosclerosis in a subject, by determining a single nucleotide polymorphism in specific codon of a polynucleotide encoding human CCR2 receptor in genome of the subject .
                                                                                                                                         MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120
                                                                                                                                                    121 HIGYFGGIFFIILLTIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTK 180
                                                                                                                                                                                                 181 CQKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR 240
                                                                                                                                                                                                                                               241 AVRVIFTIMIVYFLFWTPYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCI 300
                                                                                                                                                                                                                                                                                               7; Gaps
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                                                                                                        1 MLSTSRSRFIRNTNESGEEVTTFFDYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGN
                                                Score 1651.5; DB 22; Length 360;
Pred. No. 5.4e-180;
3; Mismatches 5; Indels 7;
                                                                       5; Indels
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human CCR2-64I polymorphic variant polypeptide.
                                                                                                                                                                                                                                                                                                                              NPIIYAFVGEKFR---SLF---HIALG-CRIAPL 327
                                                                                                                                                                                                                                                                                                                                          wild-type receptor polypeptide CCR2-64V,
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Misc-difference 64
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                                              83.8%;
95.5%;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                            (first entry)
                                                          Best Local Similarity 95.5
Matches 319; Conservative
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                        360 AA;
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        The invention relates to diagnosing atherosclerosis (or susceptibility to) in a subject by determining expression or activity of the human CCR2-64I polypeptide (a polymorphic variant form of the human CCR2 receptor) or the CCR2-64V polypeptide (human CCR2 receptor), by screening for a single nucleotide polymorphism in codon 64 of the polymucleotide encoding the CCR2 receptor. This results in production of CCR2-64I, whereby polymorphic variants are associated with a lower incidence of atherosclerosis. The presence or amount of CCR2-64I/V in a sample can also be analysed. The sequences of the invention can be used for predicting the response of a patient to draw treatment, for predicting the disease outcome in a patient and also for the production of a treatment for hypercholesterolaemia. The sequence represents the polymorphic variant polypeptide CCR2-64I.
                                                                                                                                                                                                                                                                                                                                                                                                                                                              MLVVLILINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120
                                                                                                                                                                                                                                                                                                                                                                                                   HIGYFGGIFFIILLTIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTK 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            COKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR 240
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AVRVIFTIMIVYFLEWTPYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCI 300
                                                                                                                                                                                                                                                                                                                                         Gaps
                                                                                                                                                                                                                                                                                                                      1 MLSTSRSRFIRNTNESGEEVTTFFDYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGN 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Human; G protein-coupled receptor; GPCR; non-endogenous; mutant; constitutively activated GPCR; agonist; disease.
                                                                                                                                                                                                                                                             Score 1650.5; DB 22; Length.360;
Pred. No. 7.1e-180;
4; Mismatches 5; Indels 7;
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                                                                                                                                                                                                                                                             83.8%;
95.2%;
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                                                                                                                                                                                                                                                                           Local Similarity
                                                                                                                                                                                                                               360 AA;
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Synthetic.
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Lin I;

Lehmann-Bruinsma K, Liaw CW,

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           δ
δλ
                                                                                                                                                                                                                                                                3;
                                                                                                              The invention relates to G protein-coupled receptors (GPCRs) for which the endogenous ligand has been identified. Non-endogenous constitutively activated receipens of known GPCRs are used in the invention for the direct identification of candidate compounds as
                                            use in
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    241 AKRVIFTIMIVYFLEWTPYNIVILLNTPQEFFGLSNCESTSQLDQATQVTETLGMTHCCI 300
                                                                                                                                                                                                                                                                                                                                               61 MLVVLILINCKKLKCLTDIYLLNLAISDELFLITLPIMAHSAANEWVFGNAMCKLFTGLY 120
                                                                                                                                                                                                                                                                                                                                                                               HIGYEGGIFFIILLTIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTK 180
                                                                                                                                                                                                                                                                                                                                                                                             241 AVRVIFTIMIVYFLEWTPYNIVILLNTFQEFFGLSNÇESTSQLDQATQVTETLGMTHCCI 300
                                                                                                                                                                                                                                                                                                                                 61 MLVVLILLINCKKLKCLTDIYLLNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLY 120
                                                                                                                                                                                                                                                                                                7; Gaps
                                                                                                                                                                                                                                                                                      1 MLSTSRSRFIRNTNESGEEVTTFFDYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGN 60
                                                                                                                                                          receptor agonists, inverse agonists or partial agonists. Such agonists are useful as therapeutic agents for diseases or disorders associated with GPCRs. The present sequence is a non-endogenous version of a known human GPCR.
                                                                                                                                                                                                                                                                                                                                                                                                                             181. COKEDSVYVCGPYFPRGWNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHR
                                                                                                                                                                                                                                                                                                                                                                                                                                          disease treatment, comprises contacting candidate compounds with versions of GPCRs
                                            Identifying agonists of G protein-coupled receptors (GPCRs) for
                                                                                                                                                                                                                                         DB 22; Length 360;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Human; receptor; DC-SIGN; dendritic cell; T lymphocyte; HIV;
gpl20; C-type lectin; ICAM3; HIV entry; T cell; macrophage;
HIV infection; CCR5.
                                                                                                                                                                                                                                                                6; Indels
                                                                                                                                                                                                                                         83.5%; Score 1645.5; DB 2795.2%; Pred. No. 2.6e-179; Live 3; Mismatches 6;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       NPIIYAFVGEKFR---SLF---HIALG-CRIAPL 327
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            301 NPIIYAFVGEKFRRYLSVFFRKHITKRFCKOCPV 334
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Amino acid sequence of human CCR5 protein.
                                                                                         Claim 1; Page 274-275; 394pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AAG79089 standard; Protein; 352 AA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                28-FEB-2001; 2001WO-US06322
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       02-MAR-2000; 2000US-0517605
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (first entry)
                                                                                                                                                                                                                                                      Best_Local Similarity 95.2
Matches 318; Conservative
             WPI; 2001-648759/74
                                                                                                                                                                                                                      360 AA;
                                                                    versions of GPCRs
                         N-PSDB; ABI97976
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  WO200164752-A2.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 10-DEC-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         07-SEP-2001
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAG79089;
                                                                                                                                                                                                                       Sequence
                                                                                                                                                                                                                                             Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       301
                                                                                                                                                                                                                                                                                                                                                                                  121
                                                                                                                                                                                                                                                                                                                                                                                                                                                    181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                RESULT 10
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                                                                                                                                                                                                                                                                                                                                                                                   Qγ
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lymphocytes)
                                                                                                                                                                                                                                                                                                                                                             The specification describes an antibody which is specific for an antigenic fragment of gp120. This antigenic fragment binds to DC-SIGN or is exposed upon gp120 binding of DC-SIGN due to concomitant conformational change. DC-SIGN is a receptor that is specifically expressed on dendritic cells and facilitates infection of T lymphocytes with HIV. DC-SIGN is identical to a HIV-1 gp120-binding C-type lectin. DC-SIGN binds is identical to a HIV-1 gp120-binding C-type lectin. With high affinity. The antibody of the invention inhibits the trans enhancement of HIV entry into a T cell or macrophage facilitated by
                                                                                                                                                                                               An antibody for the treatment or prevention of HIV-infection comprises a gpl20 portion which binds to DC-SIGN or is exposed upon gpl20 binding of DC-SIGN due to concomitant conformational change -
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  261 IVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCINPIIYAFVGEKFRSLF---- 316
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        145 AVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTKCQKEDSVYVCGPYFP----RGWNN 200
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLYHIGYFGGIFFIILLTIDRYLAIVH 144
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      25 DYDYGAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGNMLVVLILINCKKLKCLTDIYLLNL 84
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          FHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHRAVRVIFTIMIVYFLFWTPYN
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             dendritic cells. The antibody is useful to treat or prevent HIV infection. The present sequence represents a human CCR5 protein, which is a translocation promoting agent that interacts with CD4 This receptor functions in HIV-1 entry into cells.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  10;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Length 352;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               CC-CKR5; chemokine receptor; mouse; human; transgenic mouse; HIV infection; T-cell mediated inflammation.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        62.7%; Score 1236; DB 22;
77.4%; Pred. No. 1.8e-132;
tive 25; Mismatches 34;
                                                                                     Geijtenbeek
                                                                                                                                                                                                                                                                                                                   Disclosure; Page 118-119; 131pp; English.
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                                                                                     Van Kooyk Y,
(UYNY ) UNIV NEW YORK STATE. (UYNI-) UNIV NIJMEGEN.
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                                                                                        Kwon D,
                                                                                                                                           WPI; 2001-602565/68.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     352 AA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           317 --HIA 319
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     313 QKHIA 317
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    06-AUG-1998
                                                                                        Littman DR,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     236;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence
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usetul to diagnose, prevent and/or trea
autoimmune disease and viral infection
                                                                                                                                                                                                    Samson M,
     diagnosis; treatment; prevention;
                                                                                                                                                                                                                                                                                                Claim 4; Fig 1b-c; 94pp; English.
                                                                                                                                                        96EP-0870021,
                                                                                                                                               96EP-0870102,
                                                                                                                                                                                                  Libert F, Parmentier M,
                                                                                                                                                                             (EURO-) EUROSCREEN SA.
                                                                                                                                                                                                                      WPI; 1997-479829/44.
N-PSDB; AAT90117.
                                                                                                                                                                                                                                                                                                                                                                                                                                                 352 AA;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    310 VFFQKHIA 317
                                                                             W09732019-A2
                                                           Homo sapiens
                                                                                                                         28-FEB-1997;
                                                                                                                                              06-AUG-1996;
                                                                                                                                                         01-MAR-1996;
                                                                                                    04-SEP-1997
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                                                                                                                                                                                                                                                                                                                                                                      ä
                                                                                                                                                                                                                                  This sequence is the mouse CC-CKR5 protein, is encoded by the DNA of the Invention. CC-CKR5 is a chemokine receptor. Cells transformed with the DNA can be cultivated and the expression product harvested. The DNA can be knocked out and replaced with the human CC-CKR5 gene, creating transgenic mice which can be used in the study of HIV infection or T-cell mediated inflammation. Transgenic mice could also be used to screen for human CC-CKR5 agonists or antagonists.
                                                                                                                                                                                                                                                                                                                                                                                                                                          251 VYFLFWTPYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCINPIIYAFVGE 310
                                                                                                                                                                                                                                                                                                                                                                                                                                75 CLTDIYLLNLAISDILFLITLPLWAHSAANEWVFGNAMCKLFTGLYHIGYFGGIFFIILL 134
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       135 TIDRYLAIVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTKQQKEDSVYVCGPYF 194
                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         17 GEEVTTFFDYDYG--APCHKFDVKQIGAQLLPPLYSLVFIFGFVGNMLVVLILINCKKLK 74
                                                                                                                                                                                                                                                                                                                                                                                                 New isolated mouse chemokine receptor, CC-CKR5 - used to develop products for the study, diagnosis and treatment of HIV infection or T-cell mediated inflammation
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    195 PRG----WINFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHRAVRVIFTIMI
                                                                                                                                                                                                                                                                                                                                                                     9;
                                                                                                                                                                                                                                                                                                                                                 Length 354;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human Cys-Cys chemokine receptor 5; CCR5; human immunodeficiency virus; type 1; type 2; HIV-1; HIV-2;
                                                                                                                                                                                                                                                                                                                                                                    Indels
                                                                                                                                                                                                                                                                                                                                            62.6%; Score 1234; DB 19; 74.7%; Pred. No. 3.1e-132; ive 29; Mismatches 43;
                                                                                                                     Shabon U;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAW27407 standard; Protein; 352 AA.
                                                                                            (SMIK ) SMITHKLINE BEECHAM CORP.
                                                                                                                                                                                                               Claim 11; Fig 1; 27pp; English.
                                                                         96US-0724984
                                                   97EP-0307823
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                                                                     Local Similarity 74.79 tes 230; Conservative
                                                                                                                 Bergsma DJ, Brawner ME,
                                                                                                                                      WPI; 1998-195463/18.
                                                                                                                                                                                                                                                                                                                         354 AA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      KFRSLFHI 318
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          305 KFRSYLSV 312
                                                                                                                                                  N-PSDB; AAV23989
                                                  03-OCT-1997;
                              08-APR-1998
          EP834564-A2
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                                                                                                                                                                                                                                                                                                                         Sequence
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                                                                                                                                                                                                                                                                                                                                            Query Match
Best Local 9
                                                                                                                                                                                                                                                                                                                                                                  Matches
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82 LNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLYHIGYFGGIFFIILLTIDRYLA 141
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCINPIIYAFVGEKFRSLF- 316
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gaps
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   10 YDINXYTSEPÇQKINVKQIAARLLPPLYSLVFIFGFVGNMLVILILINCKRLKSMTDIYL 69
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            24 FDYDY -- GAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGNMLVVLILINCKKLKCLTDIYL 81
inflammatory disease; rheumatoid arthritis; glomerulonephritis; asthma; idiopathic pulmonary fibrosis; psoriasis; viral infection; cancer; atherosclerosis; autoimmune disorder.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               S (CCR5), which is stimulated by MIP-1 alpha, MIP-1 beta or RANTES chemokines, but not by monocyte chemoattractant protein 1 (MCP-1), MCP-2, MCP-3, interleukin-8 (IL-8) or growth related gene product alpha (GRO alpha) chemokines. Active CCR-5 is also a receptor of human immunodeficiency virus type 1 or type 2 (HIV-1 or HIV-2). CCR5 or its cDNA can used to diagnose, treat and/or prevent inflammatory diseases, e.g. rheumatoid arthritis, alphamatoria asthma, dialopathic pulmonary fibrosis and psoriasis, viral infections, especially HIV-1 or HIV-2 infection, cancer, atherosclerosis and autoimmune disorders.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Active and inactive forms of human CC chemokine receptor CCR-5 useful to diagnose, prevent and/or treat inflammatory disorders,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       The present sequence is human CC (Cys-Cys) chemokine receptor
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Length 4352;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       (AAT85161) isolated from a macrophage library. It shows 62% identity to CCCKR1. Chemokine receptor 88-2B (see AAW27124) has also been identified. 88C and 88-2B receptors and their polypeptide fragments
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              can be produced in transformed host cells. The receptors, peptides comprising one or more of the extracellular or intracellular domains, and anti-receptor antibodies can be used to modulate receptor activities, particularly ligand and G protein binding, are potentially potentially useful in the treatment of are potentially potentially useful in the treatment of artheroslerosis, rheumatorid arthritis, tumours, asthma, viral infection, AIDS, inflammatory conditions, pathological immune
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    New nucleic acid encoding chemokine receptors 88-2B and 88C - used
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              to modulate leukocyte trafficking, e.g. for treatment of inflammation, tumours, viral infections, autoimmune diseases, etc.
                                                                                                                                                                                                                                                                 tumour; asthma; viral infection; AIDS; inflammation;
autoimmune disease; therapy; diagnosis; leukocyte trafficking;
G protein coupled receptor; ligand; modulator; antibody; human.
                                                                                                                                                                                                                                               Chemokine receptor 88C; atherosclerosis; rheumatoid arthritis;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               response, abnormal haematopoietic processes etc.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1..32
/label= Extracellular_domain
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    89..112
/label= Extracellular_domain
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Schweickart VL;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Claim 16; Page 47-48; 65pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                           Location/Qualifiers
                          AAW27123 standard; Protein; 352 AA
                                                                                                                                                                                           Human chemokine receptor 88C.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  96WO-US20759
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 95US-0575967
                                                                                                                                         (first entry)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    .352
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Gray PW, Raport CJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 1997-341689/31.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       352 AA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (ICOS-) ICOS CORP.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  N-PSDB; AAT85161
                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         WO9722698-A2
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  20-DEC-1996;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       07-JUN-1996;
20-DEC-1995;
                                                                                                                                   14-DEC-1997
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             26-JUN-1997
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                                                                            AAW27123;
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AAW27123
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This polypeptide sequence comprises macaque chemokine receptor 88C, a G protein coupled receptor that is involved in leukocyte trafficking. Its amino sequence was deduced from a 88C DNA (AAM95163) isolated by PCK amplification. It shows 97% identity to human 88C (AAW27123). 88C receptors and their polypeptide fragments
                                                                                                                                                    198 WNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHRAVRVIFTIMIVYFLFWT 257
                                                                                                                                    82 INLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLYHIGYFGGIFFIILLTIDRYLA 141
                                                                                                                                                                                                    142 IVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTKCQKEDSVYVCGPYFP----RG 197
                                 12; Gaps
                                                                                    10 YDINYYTSEPCQKINVKQIAARILPPLYSLVFIFGFVGNMLVILILINCKRLKSMTDIYL 69
                                                                 24 FDYDY --GAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGNMLVVLILINCKKLKCLTDIYL 81
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       New nucleic acid encoding chemokine receptors 88-2B and 88C - used to modulate leukocyte trafficking, e.g. for treatment of inflammation, tumours, viral infections, autoimmune diseases, etc.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Chemokine receptor 88c; atherosclerosis; rheumatoid arthritis; tumour; asthma; viral infection; AIDS; inflammation; autoimmune disease; therapy; diagnosis; leukocyte trafficking; G protein coupled receptor; ligand; modulator; antibody.
Length 352;
                                   Indels
62.1%; Score 1224; DB 18; 76.3%; Pred. No. 4.2e-131; iive 27; Mismatches 34;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Schweickart VL;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 36; Page 57-58; 65pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAW27125 standard; Protein; 352 AA.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Macaque chemokine receptor 88C.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      96US-0661393.
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   Query Match 62.1%
Best Local Similarity 76.3%
Matches 235; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Gray PW, Raport CJ,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          WPI; 1997-341689/31.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         (ICOS-) ICOS CORP.
                                                                                                                                                                                                                                                                                                                                                                                                            317 -----HIA 319
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                                                                                                                                                                                                                                                                                                      142 IVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTKCQKEDSVYVCGPYFP----RG 197
                                                                                                                                                                                                                                                                                                                                                            258 PYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCINPIIXAFVGEKFRSLF- 316
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        198 WNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHRAVRVIFTIMIVYFLEWT 257
                                                                                                                                                                                                              Indels .12; Gaps
            comprising one or more of the extracellular or intracellular domains, and anti-receptor antibodies can be used to modulate receptor activities, particularly ligand and G protein binding, and artheroslerosis, rheumatoid arthritis, tumours, asthma, viral infection, ADS, inflammatory conditions, pathological immune response, abnormal haematopoietic processes etc. A hybridoma that produces an antibody that specifically binds to macaque 88C is
                                                                                                                                                                                                                                    24 FDYDY -- GAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGNMLVVLILINCKKLKCLTDIYL 81
                                                                                                                                                                                                                                                    can be produced in transformed host cells. The receptors, peptides
                                                                                                                                                                                     Length 352;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     CC chemokine receptor 5; CCR5; G-protein coupled receptor; human immunodeficiency virus; HIV; CD4; AIDS; therapy;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           "extracellular loop-1 (Claim 19)"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       "extracellular loop-2 (Claim 19)"
                                                                                                                                                                                62.1%; Score 1224; DB 18; 76.6%; Pred. No. 4.2e-131; ive 26; Mismatches 34;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    29.55
/label= I
/note= "transmembrane domain"
109..120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            143..171
/label- IV
/note= "transmembrane domain"
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/note= "transmembrane domain"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Human CC chemokine receptor 5 (CCR5).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         AAW23835 standard; Protein; 352 AA.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               08-JUN-1998 (first entry)
                                                                                                                                                                                             Best Local Similarity 76.69
Matches 236; Conservative
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104..126
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                                                                                                                                                    352 AA;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   transgenic animal.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             317 ----HIA 319
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        310 VFFQKHIA 317
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Homo sapiens.
                                                                                                                                                      Sequence
                                                                                                                                                                                Query Match
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     AAW23835;
                                                                                                                          claimed
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 8
                                                                                                                                                                                                                                     δλ
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This protein sequence comprises of a novel human macrophage-selective and chemokine receptor that has been designated CCR5. The sequence was deduced from an isolated cDNA clone (see AAT76920). An Ala127Leu variant (see W238340 of CCR5 was also identified. The susceptibility of human macrophages to HIV infection depends on cell surface expression of CD4 and CCR5. CCR5 is a member of the 7-transmembrane superfamily of G-protein coupled cell surface molecules. It plays an essential role in the membrane fusion step of infection by some HIV isolates. The establishment of stable, non-human coll lines and transgenic mammals having cells that coexpress human CD4 and CCR5 provides valuable tools for research of HIV infection. Antibodies that bind to CCR5, CCR5 variants, and CCR5-binding agents capable of blocking membrane fusion between HIV and target tropic strains of HIV.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           257
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              258 PYNIVILLNTFQEFFGLSNCESTSQLDQATQVTETLGMTHCCINPIIYAFVGEKFRSLF- 316
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          82 LNLAISDLLFLITLPLWAHSAANEWVFGNAMCKLFTGLYHIGYFGGIFFIILLTIDRYLA 141
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   142 IVHAVFALKARTVTFGVVTSVITWLVAVFASVPGIIFTKCQKEDSVYVCGPYFP----RG 197
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                24 FDYDY -- GAPCHKFDVKQIGAQLLPPLYSLVFIFGFVGNMLVVLILINCKKLKCLTDIYL 81
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              198 WNNFHTIMRNILGLVLPLLIMVICYSGILKTLLRCRNEKKRHRAVRVIFTIMIVYFLFWT
                                                                                                                                                                                                                                                                                                                                                                                               {\tt CC} chemokine receptor 5 polypeptide - used to inhibit membrane fusion between {\tt HIV} and a target {\tt cell}
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         62.1%; Score 1224; DB 19; Length 352; 76.3%; Pred. No. 4.2e-131; Live 27; Mismatches 34; Indels 12
                                                                                         loop-3 (Claim 19)"
                                                                                                                                                                                                                                                                                                            G, Berger EA, Broder CC, Combadiere C;
Kennedy PE, Murphy PM;
                                                         'note= "transmembrane domain"
             /note= "transmembrane domain"
                                                                                                 277..300
/label= VII
/note= "transmembrane domain"
                                                                                                                                                                                                                                                                                (USSH ) US DEPT HEALTH & HUMAN SERVICES,
                                                                                        "extracellular
                                                                                                                                                                                                                                                                                                                                                                                                                                            Claim 68; Fig 1C; 70pp; English
                                                                                                                                                                                                                        97WO-US09586
                                                                                                                                                                                                                                                      96US-0018508
                                            /label= VI
   /label = V
                                                                        ..276
                             .258
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                                                                                     /note=
                                                                                                                                                                                                                                                                                                                                                    WPI; 1998-032650/03
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            352 AA;
                                                                                                                                                                                                                                                                                                                                                                    N-PSDB; AAT76920
                                                                                                                                                                                                                     28-MAY-1997;
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                                                                                                                                                                                           04-DEC-1997.
                                                                                                                                                                                                                                                                                                          Alkhatib G,
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Query Match
                              Domain
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q
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317 ----HIA 319 ||| 310 VFFQKHIA 317 Οy

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